

IN THE NAME OF GOD

Qazvin university of medical science
Department of Immunology & Microbiology



JOURNAL CLUB & MSC SEMINAR



Present by : Sahar Amirkamali
Supervise by : Dr.M.Aslanimehr

CORONAVIRUS CLASSIFICATION

Order: <i>Nidovirales</i>	(4 Families) < history >
Family: <i>Arteriviridae</i>	(1 Genus) < history >
Family: <i>Coronaviridae</i>	(2 Subfamilies) < history >
Subfamily: <i>Coronavirinae</i>	(4 Genera) < history >
Genus: <i>Alphacoronavirus</i>	(8 Species) < history >
Genus: <i>Betacoronavirus</i>	

Species: *Betacoronavirus 1*

Species: *Human coronavirus HKU1*

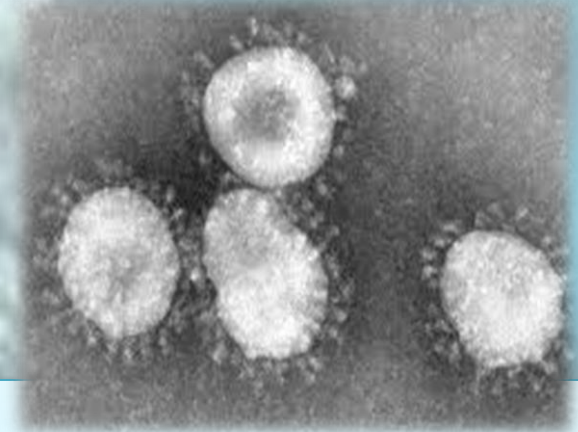
★ Species: *Murine coronavirus*

Species: *Pipistrellus bat coronavirus HKU5*

Species: *Rousettus bat coronavirus HKU9*

Species: *Human coronavirus 229E*

Species: *Human coronavirus NL63*



• Alpha

- human examples: hcov -229E, hcov -NL63
- pig, dog, and cat covs

• beta

- hcov -OC43, hcov -HKU1, hcov -**SARS**
- MHV, rat, pig and cow covs

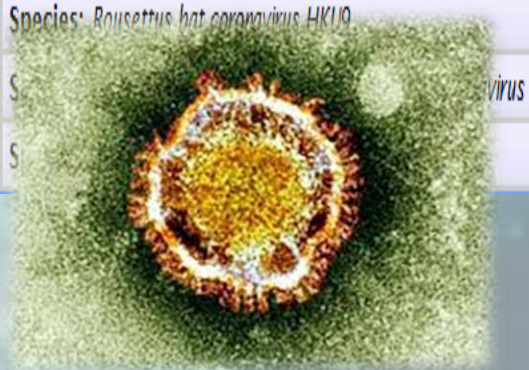
– mers-cov

• gamma

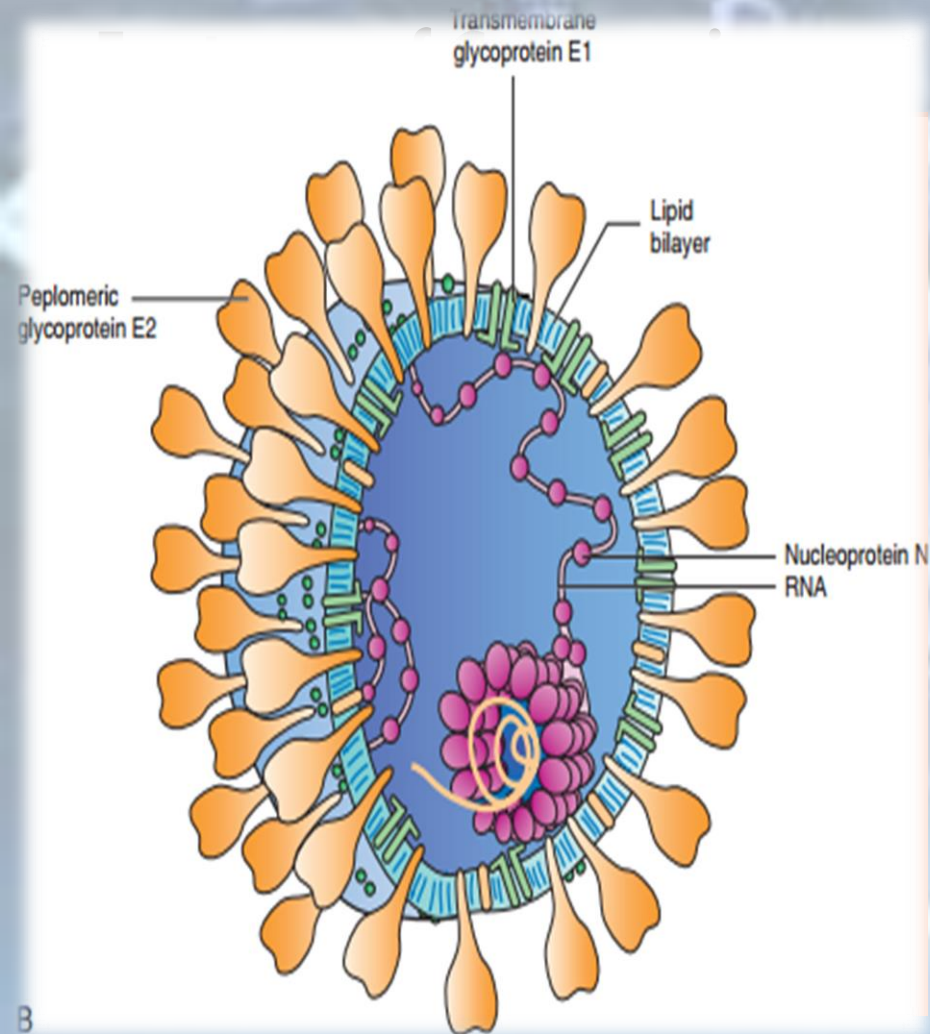
- chicken and turkey covs

• delta

- bird covs



CORONAVIRUSES (COVS):



Non-sars Human Covs: Epidemiology



Non-SARS Human CoVs: Clinical Spectrum of Illness



Severe Acute Respiratory Syndrome (SARS)



Clinical course - triphasic

SARS - morbidity

Week 1

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severe, acute
respiratory
illness



fever



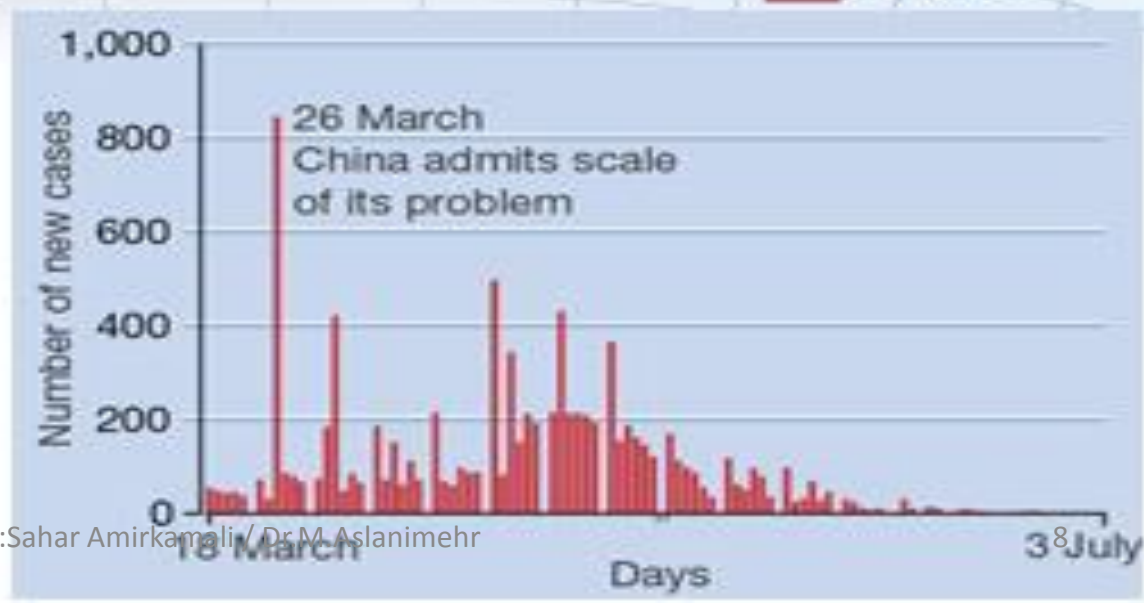
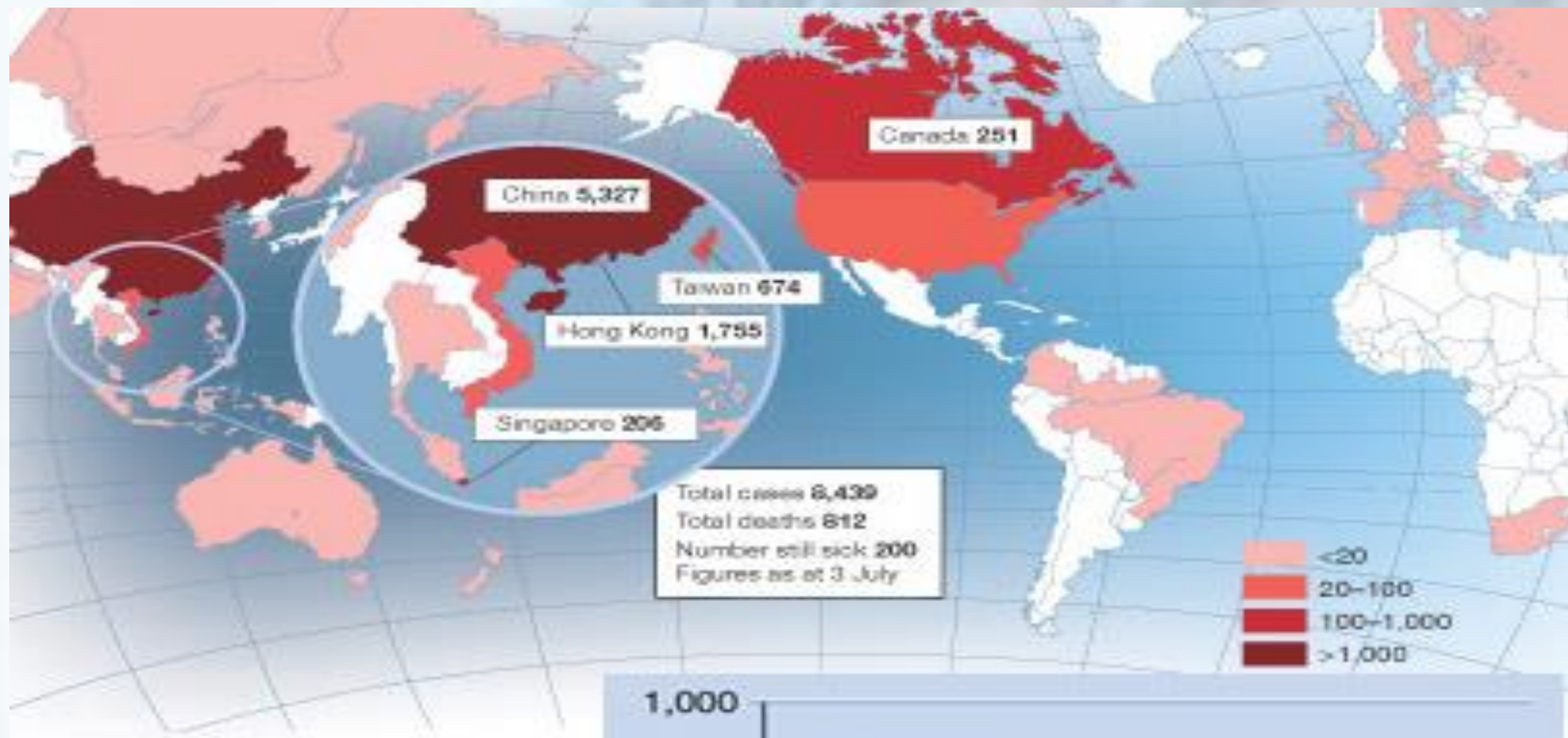
cough



breathing
difficulties

mild illness





47 cases of Middle East respiratory syndrome disease from Saudi Arabia: clinical, demographic, and clinical characteristics

Abdullah Assi
Waleh Al-Najjar

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updates from WHO, the US Centers for Disea
Control and Prevention (CDC), and the European Cent

	MERS-CoV	SARS, global ^{1,2,3,4}
Demographic factors		
Date of first case report (place)	April, 2012 (Jordan); June, 2012 (first Saudi case)	November, 2002 (China)
Mean (95% CI) incubation period (days)	5.2 (1.9–14.7); range 2–13	4.6 (3.8–5.8); range 2–14
Serial interval (days)	7.6	8.4
Age distribution	98% adults, 2% children	93% adults, 5–7% children
Mean (range) age (years)	56 (14–94)	39.9 (1–91)
Sex distribution	77% male, 23% female	43% male, 57% female
Sex ratio (male:female)	3.3:1	1:1.3
Clinical features		
Mortality	55%	0–40%
Case-fatality rate (overall)	Undefined	9.6%
In patients with comorbidities	60%	1–2%
Mean time from onset to death (days)	16.5	23.7
Presenting symptoms		
Fever >38°C	98%	99–100%
Chills or rigors	87%	15–73%
Cough	83%	62–100%
Dry cough	47%	29–75%
Productive cough	36%	4–29%
Haemoptysis	17%	0–1%
Headache	13%	20–56%
Myalgia	32%	45–61%
Malaise	38%	31–45%
Shortness of breath	72%	40–42%
Nausea	21%	20–35%
Vomiting	21%	20–35%
Diarrhoea	26%	20–25%
Sore throat	21%	13–25%
Rhinorrhoea	4%	2–24%
Comorbidities	96%	10–30%
Diabetes	68%	24%
Chronic renal disease	49%	2–6%
Chronic heart disease	28%	10%
Malignant disease	2%	3%
Hypertension	34%	19%
Obesity	17%	–
Smoking	23%	17%
Viral hepatitis	Not known	27%
Ventilatory support needed	80%	14–20%

	Female	Male	All
Dead	Total	Dead	
0	2	0	
1	1	1	
2	5	2	
4	10	4	
5	9	6	
4	11	8	
4	5	4	
1	3	2	
1	1	1	
12 (61%)	47	28 (60%)	

le East respiratory syndrome

Myalgia	30%	14–20% (30%)
Headache	13%	6 (13%)

Table 3. Symptoms of Middle East respiratory syndrome in 47 Saudi cases at presentation

By: Sahar Amirkamali / Dr M Aslanimehr

Middle East Respiratory Syndrome

BACKGROUND

- First identified in September **2012**
- Cases retrospectively identified as early as March 2012
- Different from other coronaviruses in humans, including SARS
- Most similar to coronaviruses found in bats



Hospital Day 1



Hospital Day 3

N Engl J Med 367;19, Nov 2012

MERS-COV

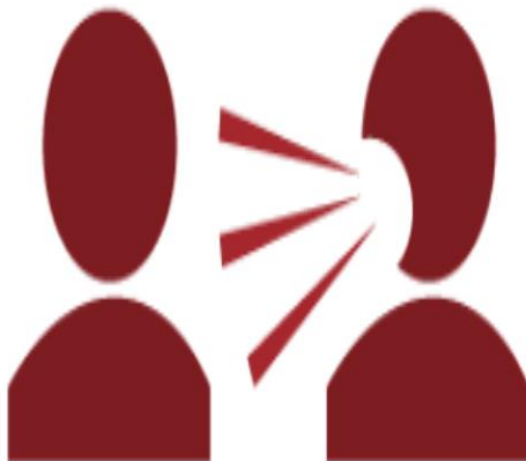
Origins



TRANSMISSION

The World Health Organization has said it is not yet known how people are infected with the virus. Investigations are being conducted to determine the source of the virus, the types of exposure that lead to infection, the mode of transmission, and the clinical pattern and course of disease.

The source of MERS has yet to be determined although camels have been identified as possible carriers



Human-to-human transmission has been recorded in healthcare facilities, among family members and between co-workers. But the mechanism by which transmission occurred, whether respiratory, direct physical contact with a patient or contamination of the environment, is still unknown.

MERS-C

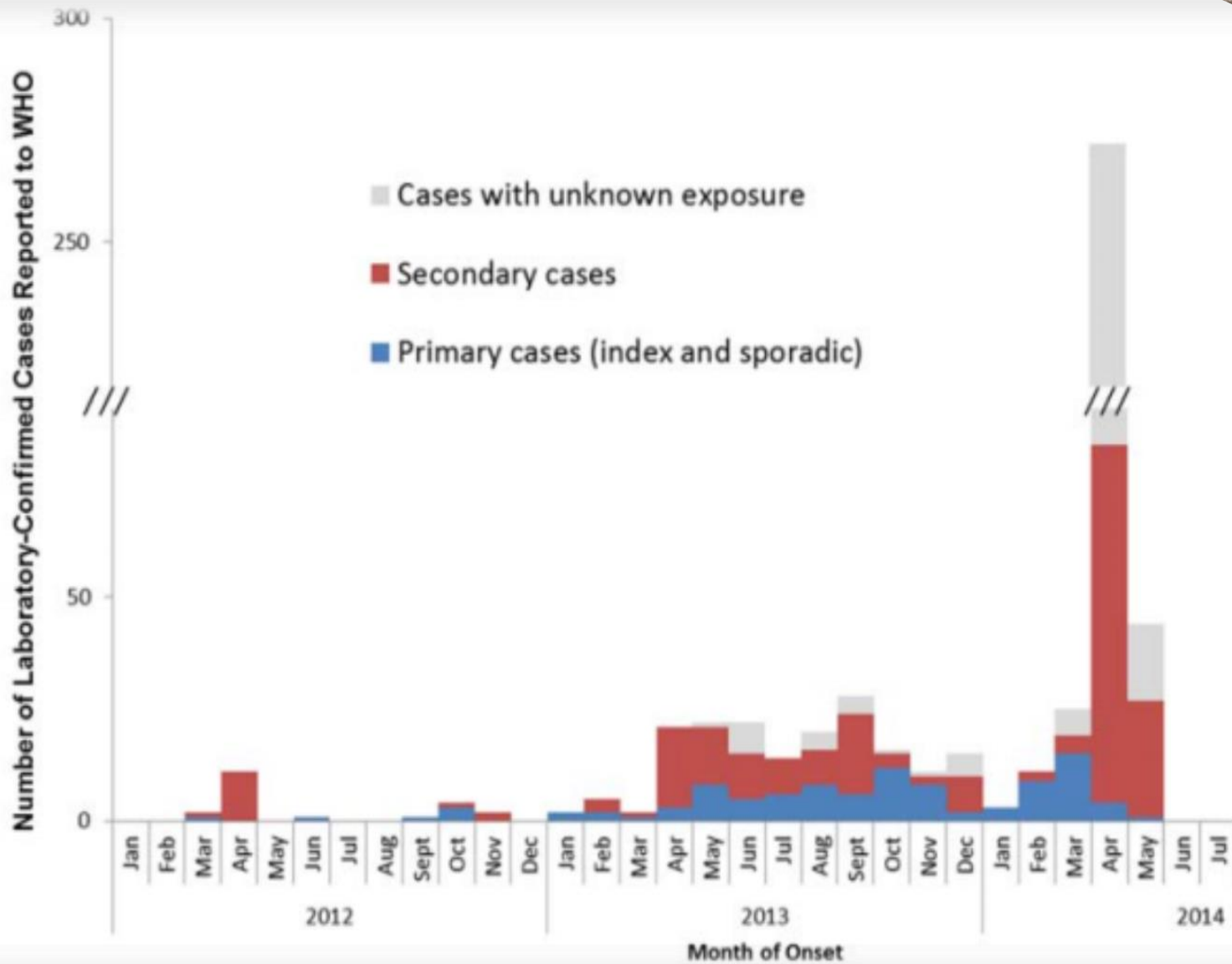
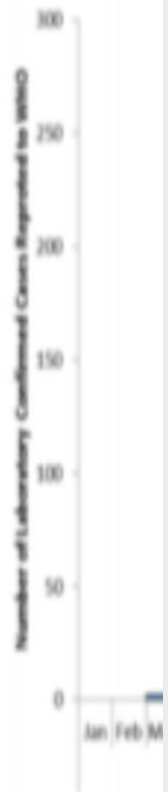


Figure 2. Epidemic curve of MERS-CoV (May 2014)

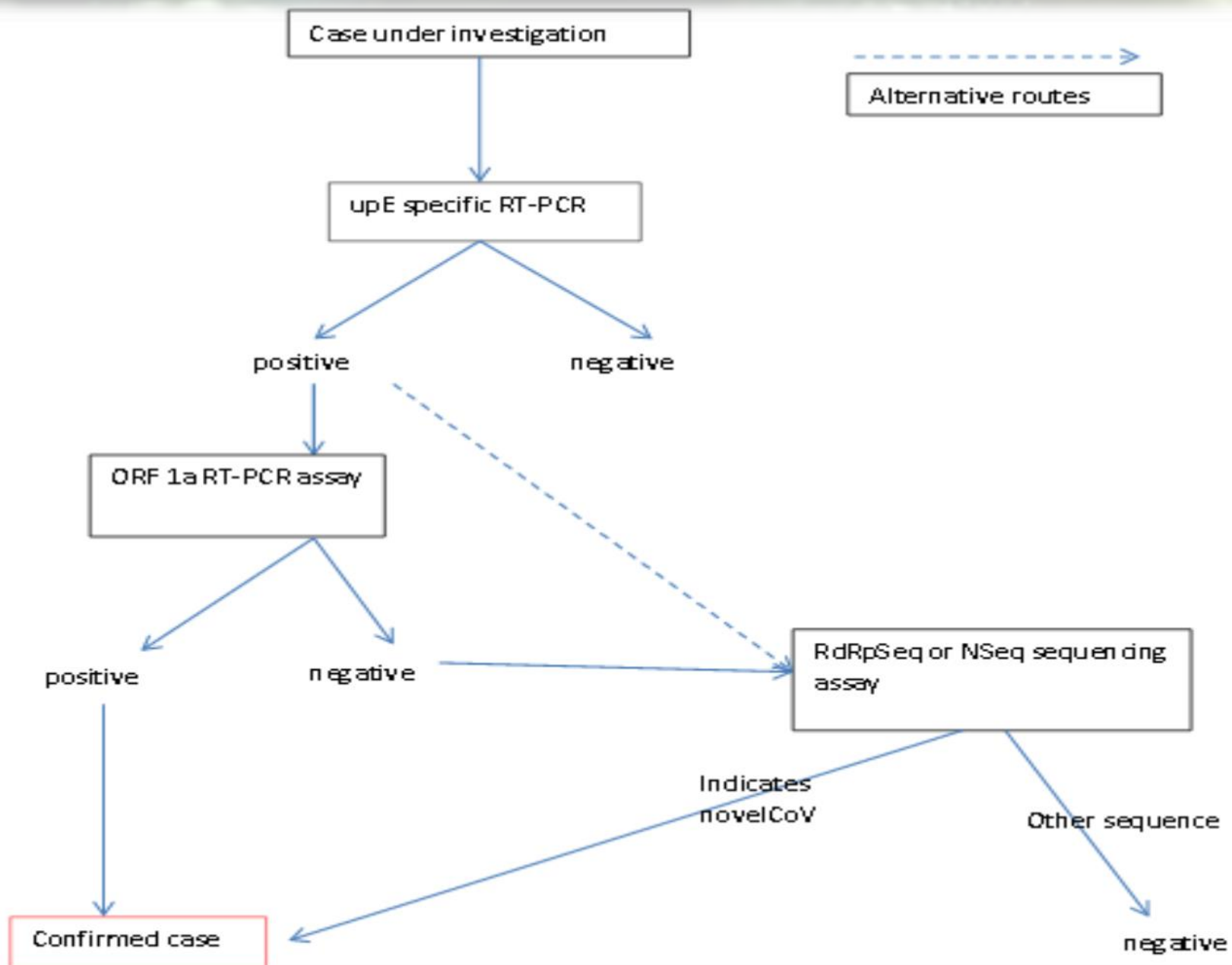
Infection Control



close contacts

Environmental Infection Control





PRECAUTIONS AGAINST MERS



1. Frequently wash your hands with soap or hand rubs with alcohol.



5. If you are a health worker, strictly follow infection control protocols in your work.



2. Practice proper cough etiquette by covering your mouth and nose while sneezing or coughing. Use facial tissues and dispose in a waste basket.



6. Visit your doctor, a hospital or health facility immediately if symptoms of MERS-CoV manifest, including persistent coughing.



3. Avoid contact with farm and domesticated animals, including camels.



7. If you were in close contact with a confirmed MERS-CoV patient, comply with local health regulations and postpone any trip abroad until after test results are negative.



4. Avoid contact with the sick or those infected with MERS-CoV. If you have respiratory illness, stay home and wear a surgical mask to protect your family.



8. Get regular exercise, eat a balanced diet, and get adequate sleep as all these will help strengthen the body's immunity.

provide insight into bat-to-human transmission of MERS coronavirus

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Edited by Michael Farzan, The Scripps Research Institute, Jupiter, Florida, and accepted by the Editorial Board, 2014

Middle East respiratory syndrome coronavirus (MERS-CoV) currently spreads in humans and causes ~36% fatality in infected patients. Believed to have originated from bats, MERS-CoV is genetically related to bat coronaviruses HKU4 and HKU5. To understand how bat coronaviruses transmit to humans, we investigated the receptor usage and cell entry activity of the virus-surface spike proteins of HKU4 and HKU5. We found that dipeptidyl peptidase 4 (DPP4), the receptor for MERS-CoV, is also the receptor for HKU4, but not HKU5. Despite sharing a common receptor, MERS-CoV and HKU4 spikes demonstrated functional differences. First, whereas MERS-CoV prefers human DPP4, HKU4 shows the opposite preference. Second, in the absence of exogenous proteases, both MERS-CoV and HKU4 spikes mediate pseudotyped virus entry into bat cells, but not human cells. Thus, HKU4 spikes mediate pseudotyped MERS-CoV spike, but not HKU4 spike, mediate pseudotyped MERS-CoV, but not HKU4 human cellular proteases. These findings suggest that DPP4 recognition is critical for cross-species transmission of MERS-CoV and HKU4.

As of June 16, 2014, Middle East respiratory syndrome coronavirus (MERS-CoV) has infected 700 people, with a fatality rate of 36% (1). MERS-CoV is a novel coronavirus that has caused human-to-human transmission studies have been limited to animals, with bats as the likely intermediate hosts (2). MERS-CoV or through camels, pose a threat to human health. Phylogenetic analysis indicates that MERS-CoV is genetically related to bat coronaviruses HKU4 and HKU5 (3-5). Understanding the transmission of these long-term emerging diseases is critical for controlling the spread of MERS-CoV. This study investigates the receptor usage and cell entry mechanism of HKU4 compared with MERS-CoV. Our results reveal that MERS-CoV has adapted to use human receptor and cellular proteases for efficient human cell entry, whereas HKU4 can potentially follow-up and also infect human cells. These findings are critical for evaluating emerging disease potentials of bat coronaviruses for preventing and controlling their spread in humans.

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Significance

A constant and long-term threat to human health is cross-species transmission of Middle East respiratory syndrome coronavirus (MERS-CoV) from bats to humans. However, this process is poorly understood. Examining the cross-species transmissibility of bat coronavirus HKU4, which is genetically related to MERS-CoV, can provide critical information about the likely causes of MERS-CoV infections in humans. Here we investigate the receptor usage and cell entry mechanism of HKU4 compared with MERS-CoV. Our results reveal that MERS-CoV has adapted to use human receptor and cellular proteases for efficient human cell entry, whereas HKU4 can potentially follow-up and also infect human cells. These findings are critical for evaluating emerging disease potentials of bat coronaviruses for preventing and controlling their spread in humans.

By: Sahar Amirizadeh / Dr. M. Aslani

Related Studies

- 2014 ,Jean Kaoru Millet and Gary R. Whittaker¹, Host cell entry of Middle East respiratory syndrome coronavirus after two-step, furin-mediated activation of the spike protein.

2014, Yang Yang, Receptor usage and cell entry of bat coronavirus HKU4 provide insight into bat-to-human transmission of MERS coronavirus

Assiri A, MERS-CoV Investigation Team. Hospital Outbreak of Middle East Respiratory Syndrome Coronavirus. N Engl J Med. 2013 Jun 19. [Epub ahead of print].

Guery B, the MERS-CoV study group. Clinical features and viral diagnosis of two cases of infection with Middle East Respiratory Syndrome coronavirus: a report of nosocomial transmission. Lancet. 2013 May 30.

<http://emergency.cdc.gov/coca>



THANK
YOU

